Rooke 10/626,571

21/12/2004

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L38 ANSWER 1 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2003:950040 HCAPLUS

DOCUMENT NUMBER: 140:19764

TITLE: Methods of inducing the expression of bone

morphogenetic proteins (BMPs) and transforming growth

factor-beta proteins (TGF- β s) in cells

INVENTOR(S): Mckay, William F.; Boden, Scott D.; Yoon,

Sangwook T.

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 81 pp., Cont.-in-part of U.S.

Ser. No. 292,951.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE		
US 2003225021	A1	20031204	US 2003-382844	-	20030307		
US 2003180266	A1	20030925	US 2002-292951		20021113		
PRIORITY APPLN. INFO.:			US 2001-331321P	P	20011114		
			US 2002-292951	A2	20021113		
			US 1988-124238	Α	19880729		
		,	US 2000-959578	Δ	20000428		

- AB A method of inducing the expression of one or more bone morphogenetic proteins and/or transforming growth factor- β proteins in a cell is described. The method includes transfecting a cell with an isolated nucleic acid comprising a nucleotide sequence encoding a LIM mineralization protein operably linked to a promoter. The one or more bone morphogenetic proteins can be BMP-2, BMP-4, BMP-6, BMP-7 or combinations thereof. The transforming growth factor- β protein can be transforming growth factor- β 1 protein (TGF- β 1). Transfection may be accomplished ex vivo or in vivo by direct injection of virus or naked DNA, or by a nonviral vector such as a plasmid. The method can be used to induce bone formation in osseous cells or to stimulate proteoglycan and/or collagen production in cells capable of producing proteoglycyan and/or collagen (e.g., intervertebral disk cells).
- IC ICM A61K048-00
 - ICS C12N005-08; C12N015-861; C12N015-867
- NCL 514044000; 424093200; 435456000; 435366000
- CC 63-1 (Pharmaceuticals)
 - Section cross-reference(s): 3, 6, 14
- ST bone morphogenetic protein transforming growth factor gene therapy; BMP TGF gene bone formation intervertebral disk disease; oligonucleotide transformation cell implant bone disease
- IT Bone morphogenetic proteins
 - RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (2; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)
- IT Bone morphogenetic proteins
 - RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 - (4; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in

21/12/2004

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cells)
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IT Bone morphogenetic proteins

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(6; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Bone morphogenetic proteins

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(7; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Adenoviridae

(AdLMP-1; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Gene, animal

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(BMP-2; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Gene, animal

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(BMP-4; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Gene, animal

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(BMP-6; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Gene, animal

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(BMP-7; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Proteins

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(HLMP-1; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Proteins

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(HLMP-1s; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Proteins

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(HLMP-2; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT Proteins

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RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (HLMP-3; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
     Proteins
TТ
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (LIM domain-containing; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
IT
     Proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (LMP-1; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-βs) in cells)
IT
     Proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (RLMP; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
     Gene, animal
IT
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (TGF-β1; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
IT
     Cell
        (annulus fibrosus; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-βs) in cells)
IT
     Spinal column, disease
        (intervertebral disk hernia; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
IT
     Spinal column
        (intervertebral disk; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
IT
     Stem cell
        (mesenchymal; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-βs) in cells)
IT
     Adenoviral vectors
     Bone formation
     Gene therapy
     Genetic vectors
     Hematopoietic precursor cell
     Mammalia
     Mesenchyme
     Molecular cloning
     Nucleic acid hybridization
     Ore genesis
     Plasmid vectors
     Retroviral vectors
     Retroviridae
     Transformation, genetic
     Transplant and Transplantation
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Viral vectors
     Virus
        (methods of inducing the expression of bone morphogenetic proteins
        (BMPs) and transforming growth factor-beta proteins (TGF-βs) in
     Oligonucleotides
ΙT
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (methods of inducing the expression of bone morphogenetic proteins
        (BMPs) and transforming growth factor-beta proteins (TGF-βs) in
        cells)
IT
     Bone morphogenetic proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (methods of inducing the expression of bone morphogenetic proteins
        (BMPs) and transforming growth factor-beta proteins (TGF-\betas) in
        cells)
     Nucleic acids
TT
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (methods of inducing the expression of bone morphogenetic proteins
        (BMPs) and transforming growth factor-beta proteins (TGF-\betas) in
        cells)
IT
     Promoter (genetic element)
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (methods of inducing the expression of bone morphogenetic proteins
        (BMPs) and transforming growth factor-beta proteins (TGF-\betas) in
        cells)
IT
     Cytomegalovirus
        (promoter; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
     Cell nucleus
IT
        (pulposus; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
IT
     Animal cell
        (somatic; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
IT
     Transforming growth factors
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (\beta-; methods of inducing the expression of bone morphogenetic
        proteins (BMPs) and transforming growth factor-beta proteins
        (TGF-\beta s) in cells)
IT
     630143-01-0
                   630143-02-1
     RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (oligonucleotide; methods of inducing the expression of bone
        morphogenetic proteins (BMPs) and transforming growth factor-beta
        proteins (TGF-\betas) in cells)
IT
     630150-76-4
                   630150-77-5
                                  630150-78-6
                                                630150-79-7
                                                              630150-80-0
     630150-81-1
                   630150-82-2
                                  630150-83-3
                                                630150-85-5
                                                              630150-86-6
     630150-87-7
                   630150-88-8
                                  630150-89-9
                                                630150-90-2
                                                              630150-91-3
     630150-92-4
                   630150-93-5
                                  630150-94-6
                                                630150-95-7
                                                              630150-96-8
     630150-97-9
                   630150-98-0
                                 630150-99-1
                                                630151-00-7
                                                              630151-01-8
     630151-02-9
                   630151-03-0
                                  630151-04-1
                                                630151-06-3
                                                              630151-07-4
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630151-13-2

630151-12-1

630151-08-5

630151-10-9

RL: PRP (Properties)

(unclaimed nucleotide sequence; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT 630150-75-3 630150-84-4 630151-05-2 630151-09-6 630151-11-0 RL: PRP (Properties)

(unclaimed protein sequence; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

IT 630151-14-3 630151-15-4 630151-16-5 630151-17-6 630151-18-7 630151-19-8 630151-20-1 630151-21-2 630151-22-3

RL: PRP (Properties)

(unclaimed sequence; methods of inducing the expression of bone morphogenetic proteins (BMPs) and transforming growth factor-beta proteins (TGF- β s) in cells)

L38 ANSWER 2 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2003:397004 HCAPLUS

DOCUMENT NUMBER: 138:397329

TITLE: cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk

degeneration and disk injury

INVENTOR(S): McKay, William F.; Boden, Scott D.; Yoon,

Sangwook T.

PATENT ASSIGNEE(S): Medtronic Sofamor Danek, USA

SOURCE: PCT Int. Appl., 94 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2.

PATENT INFORMATION:

PA	TENT	KIN	DATE		i	APPL	ICAT		DATE									
	O 2003042368					A2 20030522				WO 2	002-1		20021114					
WO																		
	W:	ΑE,																
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	
							IN,											
							MD,											
					•		SD,		•	•	•	•	•	•		•	•	
			-	•	•		VC,		•	•			10,	,	,	,	,	
	₽W•	GH,											2M	7 W	λM	A 7	RV	
	100.																	
			-	-			TM,				•	•		•		•	•	
							IT,								BF,	ВJ,	CF,	
		CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MŔ,	NΕ,	SN,	TD,	TG				
US	2003	1802	66		A1 20030925 US 2002-292951									20021113				
EP	1465	489			A2	A2 20041013 EP 2002-780657								20021114				
	R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,	
		ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	SK			
PRIORIT	Y APP												P 20011114					
									US 2002-292951						A 2	0021	113	
									τ	JS 1	988-	1242	38	7	1	9880	729	
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AB Me	thods	of	evnre	ecci	na I.	TM m	iner	alig:										

AB Methods of expressing LIM mineralization protein in non-osseous mammalian cells, such as stem cells or intervertebral disk cells (e.g., cells of the annulus fibrosus, or cells of the nucleus pulposus) are described. The methods involve transfecting the cells with an isolated nucleic acid comprising a nucleotide sequence encoding a LIM mineralization protein

operably linked to a promoter. Transfection may be accomplished ex vivo or in vivo by direct injection of virus or naked DNA, or by a nonviral vector such as a plasmid. Expression of the LIM mineralization protein can stimulate proteoglycan and/or collagen production in cells capable of producing proteoglycan and/or collagen. Methods for treating disk disease associated with trauma or disk degeneration are also described.

- IC ICM C12N
- CC 3-3 (Biochemical Genetics)
 Section cross-reference(s): 1, 6, 13
- ST cDNA LIM mineralization protein human rat sequence; disk degeneration injury therapy LMP protein splicing isoform
- IT Bone morphogenetic proteins
 - RL: BSU (Biological study, unclassified); BIOL (Biological study)
 (2, LMP protein in stimulating synthesis of; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)
- IT Adenoviral vectors

(AdHLMP-1, LIM mineralization protein cDNA cloning in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Protein motifs

(LIM domain, in LMP proteins; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Plasmid vectors

Retroviral vectors

Viral vectors

(LIM mineralization protein cDNA cloning in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

- IT Proteoglycans, biological studies
 - RL: BSU (Biological study, unclassified); BIOL (Biological study)
 (LIM mineralization protein in stimulating synthesis of; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)
- IT mRNA

RL: ANT (Analyte); BSU (Biological study, unclassified); ANST (Analytical study); BIOL (Biological study)

(LIM mineralization protein; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT cDNA

RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (LIM mineralization protein; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Proteins

RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (LMP (LIM mineralization protein), isoforms 1,2 and 3, of rat and human; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Bone morphogenetic proteins

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(LMP protein in inducing synthesis of; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Osteocalcins

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(LMP protein in stimulating secretion of; cDNAs encoding rat and human
LIM mineralization proteins and their use in treatment of disk
degeneration and disk injury)

IT Collagens, biological studies

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(LMP protein in stimulating synthesis of, as carrier for LMP protein implant in vertebral disk; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Aggrecans

RL: BSU (Biological study, unclassified); BIOL (Biological study)
(LMP protein in stimulating synthesis of; cDNAs encoding rat and human
LIM mineralization proteins and their use in treatment of disk
degeneration and disk injury)

IT Bone formation

(LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT RNA splicing

(LMP protein mRNA; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Body, anatomical

(back, disease, pain, lower; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Pain

(back, lower; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Animal cell

Gene therapy

Human

Mammalia

Nucleic acid hybridization

Rattus

(cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Promoter (genetic element)

RL: BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(cytomegalovirus, for LMP proteins synthesis; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Bone, disease

(degenerative disk disease, spine stenosis; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Probes (nucleic acid)

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(for LMP protein cDNA; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Primers (nucleic acid)

RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); PRP (Properties); ANST (Analytical study); BIOL (Biological study); USES (Uses)

(for LMP protein cDNA; cDNAs encoding rat and human LIM mineralization

proteins and their use in treatment of disk degeneration and disk injury)

IT cDNA sequences

(for LMP proteins of human and rat; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Drug delivery systems

(implants, LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Drug delivery systems

(injections, LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Spinal column, disease

(intervertebral disk hernia; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Spinal column

(intervertebral disk; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Spinal cord

(lumbar, fusion, LMP protein in gene therapy in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Bone formation

(mineralization, LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Molecular cloning

(of LIM mineralization proteins; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Protein sequences

(of LMP proteins of human and rat; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Cell differentiation

(osteoblast, LMP protein in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Stem cell

(pluripotent, LIM mineralization protein mRNA in; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Cytomegalovirus

(promoter for LMP proteins synthesis; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Mutation

(splice site, LMP protein; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Glycosaminoglycans, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study) (sulfated, LMP protein in inducing synthesis of; cDNAs encoding rat and human LIM mineralization proteins and their use in treatment of disk degeneration and disk injury)

IT Polymers, biological studies

IT

ΙT

IT

IT

IT

IT

IT

IT

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RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological
study); USES (Uses)
   (synthetic, as carrier for LMP protein containing cell used in
   intervertebral disk implant; cDNAs encoding rat and human LIM
   mineralization proteins and their use in treatment of disk degeneration
   and disk injury)
Spinal column
   (vertebra, annulus fibrosus, nucleus pulposus, LIM mineralization
   protein mRNA in; cDNAs encoding rat and human LIM mineralization
   proteins and their use in treatment of disk degeneration and disk
   injury)
             530167-59-0
530167-58-9
                           530167-62-5
                                         530167-64-7
RL: BSU (Biological study, unclassified); PRP (Properties); THU
(Therapeutic use); BIOL (Biological study); USES (Uses)
   (amino acid sequence; cDNAs encoding rat and human LIM mineralization
   proteins and their use in treatment of disk degeneration and disk
   injury)
256606-43-6, GenBank AC023788
RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
(Biological study)
   (cDNAs encoding rat and human LIM mineralization proteins and their use
   in treatment of disk degeneration and disk injury)
530167-57-8
             530167-60-3
                           530167-61-4
                                         530167-63-6
RL: BSU (Biological study, unclassified); PRP (Properties); THU
(Therapeutic use); BIOL (Biological study); USES (Uses)
   (nucleotide sequence; cDNAs encoding rat and human LIM mineralization
   proteins and their use in treatment of disk degeneration and disk
   injury)
530167-30-7
             530167-31-8
RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); PRP
(Properties); ANST (Analytical study); BIOL (Biological study); USES
(Uses)
   (primer sequence; cDNAs encoding rat and human LIM mineralization
   proteins and their use in treatment of disk degeneration and disk
   injury)
530170-01-5, 3: PN: WO03042368 SEQID: 3 unclaimed DNA
                                                       530170-02-6
530170-03-7 530170-04-8, 6: PN: W003042368 SEQID: 6 unclaimed DNA
530170-05-9, 7: PN: WO03042368 SEQID: 7 unclaimed DNA 530170-06-0, 8:
PN: WO03042368 SEQID: 8 unclaimed DNA 530170-07-1 530170-08-2
             530170-10-6
530170-09-3
                           530170-11-7
                                        530170-12-8 530170-13-9
530170-14-0 530170-15-1
                           530170-18-4
530170-19-5 530170-20-8
                           530170-21-9
                                        530170-22-0
                                                       530170-23-1
530170-24-2
             530170-25-3
                           530170-26-4
                                         530170-28-6
                                                       530170-29-7
             530170-31-1
530170-30-0
                           530170-33-3
                                         530170-34-4
                                                       530170-35-5
530170-36-6
             530170-37-7 530170-38-8 530170-39-9 530170-40-2
RL: PRP (Properties)
   (unclaimed nucleotide sequence; cDNAs encoding rat and human LIM
   mineralization proteins and their use in treatment of disk degeneration
  and disk injury)
530170-27-5
RL: PRP (Properties)
   (unclaimed protein sequence; cDNAs encoding rat and human LIM
   mineralization proteins and their use in treatment of disk degeneration
   and disk injury)
530158-96-4
RL: PRP (Properties)
   (unclaimed sequence; cDNAs encoding rat and human LIM mineralization
  proteins and their use in treatment of disk degeneration and disk
  injury)
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L38 ANSWER 3 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2000:553459 HCAPLUS

DOCUMENT NUMBER: 133:155511

TITLE: Highly-mineralized osteogenic sponge compositions, and

uses thereof

INVENTOR(S): McKay, William F.

PATENT ASSIGNEE(S): SDGI Holdings, Inc., USA SOURCE: PCT Int. Appl., 34 pp.

CODEN: PIXXD2

CODEN: PIXXL

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

P.	ATENT	KIN	DATE				LICAT		DATE										
WC	WO 2000045871					A1 20000							2000020						
											BR,								
		-		_	-				•		GE,	•					-		
											LK,								
		MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL	PT,	RO,	RU,	SD,	SE,	SG,	SI,		
		SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	UG,	us,	UZ,	VN,	YU,	ZA,	ZW,	AM,		
				•	ΚZ,	•	•	•											
•	RW:										UG,								
		-						•			MC,			SE,	BF,	ВJ,	CF,		
		-	-	-	-	•					SN,				_				
														20000204					
						1 20011107 EP 2000-905989							89	20000204					
E	1150																		
	R:							FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,		
		ΙE,	SI,	LT,	LV,	FI,	RO												
JI	2002	25360	77		T2		2002	1029		JP 2	2000-!	5969:	90	20000204					
ΑT	2533	885			E		2003	1115		AT 2000-905989					20000204				
ΑU	7726	82			B2		2004	0506		AU 2	2000-2	2756	8		2	0000	204		
ES	2209	820			Т3		2004	0701		ES 2	2000-9	9059	89		2	0000	204		
PRIORIT	Y API	PLN.	INFO	. :						US 1	1999-	1186	15P]	P 1	9990	204		
										WO 2	7-000	JS30	7	W 20000204					

- AB Osteogenic sponge compns. having enhanced osteoinductive properties for use in bone repair are described. The compns. include a quickly resorbable porous carrier, a more slowly resorbed mineral scaffold and an osteogenic factor, preferably a bone morphogenetic protein. The compns. enable increased osteoinductive activity while retaining a reliable scaffold for the formation of new bone at an implant site. Methods for therapeutic use of the compns. are also described.
- IC ICM A61L027-22
 - ICS A61L027-56; A61L027-46; A61K038-18
- CC 63-7 (Pharmaceuticals)
 - Section cross-reference(s): 2
- ST osteogenic sponge morphogenetic protein bone implant
- IT Bone morphogenetic proteins
 - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
- (2; highly-mineralized osteogenic sponge compns. for repair of bone)
 IT Bone morphogenetic proteins
 - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(7; highly-mineralized osteogenic sponge compns. for repair of bone) Proteins, specific or class IT RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (LMP (LIM-mineralization proteins); highly-mineralized osteogenic sponge compns. for repair of bone) IT Ceramics (biocompatible; highly-mineralized osteogenic sponge compns. for repair of bone) Bone formation ΙT Osteoblast Osteoclast Particle size distribution (highly-mineralized osteogenic sponge compns. for repair of bone) IT Bone morphogenetic proteins Collagens, biological studies Platelet-derived growth factors Steroids, biological studies RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (highly-mineralized osteogenic sponge compns. for repair of bone) IT (implant; highly-mineralized osteogenic sponge compns. for repair of bone) ΙT Porosity (microporosity; highly-mineralized osteogenic sponge compns. for repair of bone) IT Bone marrow (osteogenic enhancing factor of; highly-mineralized osteogenic sponge compns. for repair of bone) IT Growth factors, animal RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (osteogenins; highly-mineralized osteogenic sponge compns. for repair of bone) IT Bone (particles of; highly-mineralized osteogenic sponge compns. for repair of bone) IT Surgery (spinal fusion; highly-mineralized osteogenic sponge compns. for repair of bone) IT Medical goods (sponges, osteogenic; highly-mineralized osteogenic sponge compns. for repair of bone) IT Spinal column (vertebra, fusion of; highly-mineralized osteogenic sponge compns. for repair of bone) TΤ Transforming growth factors RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (β-; highly-mineralized osteogenic sponge compns. for repair of bone)

IT

Microglobulins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

 $(\beta\text{-microglobulins}; \text{ highly-mineralized osteogenic sponge compns.}$ for repair of bone)

IT 10103-46-5, Calcium phosphate

RL: DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(biocompatible ceramics; highly-mineralized osteogenic sponge compns. for repair of bone)

IT 61912-98-9, Insulin like growth factor 62031-54-3, Fgf
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); DEV (Device component use); PEP (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(highly-mineralized osteogenic sponge compns. for repair of bone)

REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L38 ANSWER 4 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2000:553458 HCAPLUS

DOCUMENT NUMBER: 133:155510

TITLE: Osteogenic paste compositions and uses thereof

INVENTOR(S): McKay, William F.

PATENT ASSIGNEE(S): SDGI Holdings, Inc., USA SOURCE: PCT Int. Appl., 37 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.						KIND DATE APPLICATION NO.												
	WO	2000	A1	2000	0810					20000204									
		W:	ΑE,	AL,	AM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CR,	CU,	
								ES,											
								KP,											
								MX,											
								TT,											
								RU,			·	•	·	•	•	•	•	•	
		RW:	GH,	GM,	KE,	LS,	MW,	SD,	SL,	SZ,	TZ,	UG,	ZW,	AT,	BE,	CH,	CY,	DE.	
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						LV,					-	•	•	•	•	•	•	•	
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	AU 770196							2004	0212		AU 2	000-2	27564	4	20000204				
	US 2004002558							2004	0101	1	US 2	001-	9231	17	20010806				
PRIORITY APPLN. INFO.:										1	US 1	999-:	1186	14P	I	2 19	9902	204	
										1	WO 2	000-1	JS302	24	V	V 2	00002	204	
AB	Des	scrib	ed a:	re o	steo	geni	с ра	ste (compi	ns. v	with	enha	anced	d ost	eoir	duct	ive		

AB Described are osteogenic paste compns. with enhanced osteoinductive properties for use in bone repair. Compns. comprising a quickly resorbable paste carrier, a more slowly resorbed mineral matrix, and Bone Morphogenetic Protein (BMP) or other osteogenic factor are described which enable increased osteoinductive activity while retaining a reliable

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scaffold for the formation of new bone at the implant site. Methods for
     making and methods for therapeutic use of the compns. are also disclosed.
IC
     ICM A61L027-22
     ICS A61L027-46; A61K038-18; A61L027-58
     63-7 (Pharmaceuticals)
CC
ST
     osteogenic bone paste implant
ΙT
     Bone morphogenetic proteins
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (2; osteogenic paste compns. and uses thereof)
TΤ
     Bone morphogenetic proteins
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (4; osteogenic paste compns. and uses thereof)
     Bone morphogenetic proteins
IT
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (6; osteogenic paste compns. and uses thereof)
TΤ
     Bone morphogenetic proteins
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (7; osteogenic paste compns. and uses thereof)
TT
     Proteins, specific or class
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (LMP (LIM-mineralization protein); osteogenic paste compns. and uses
        thereof)
IT
     Prosthetic materials and Prosthetics
        (bioactive glass; osteogenic paste compns. and uses thereof)
TT
     Polymers, biological studies
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (biocompatible non-resorbable; osteogenic paste compns. and uses
        thereof)
IT
     Ceramics
        (biocompatible; osteogenic paste compns. and uses thereof)
IT
        (demineralized; osteogenic paste compns. and uses thereof)
IT
     Coral
        (hydroxyapatite of; osteogenic paste compns. and uses thereof)
IT
        (implant; osteogenic paste compns. and uses thereof)
IT
     Anti-inflammatory agents
        (nonsteroidal; osteogenic paste compns. and uses thereof)
IT
    Antibiotics
     Body temperature
    Bone formation
    Bone marrow
     Fungicides
    Osteoblast
    Osteoclast
    Particle size distribution
    Spinal column
    Wetting agents
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(osteogenic paste compns. and uses thereof)
     Gelatins, biological studies
ΙT
     Platelet-derived growth factors
     Steroids, biological studies
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (osteogenic paste compns. and uses thereof)
IT
     Growth factors, animal
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PEP (Physical, engineering or chemical process); THU
     (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
        (osteogenins; osteogenic paste compns. and uses thereof)
     Bone marrow
IT
        (osteoprogenitor cell; osteogenic paste compns. and uses thereof)
     Drug delivery systems
TΤ
     Medical goods
        (pastes; osteogenic paste compns. and uses thereof)
     Spinal column
TT
        (vertebra, fusion of; osteogenic paste compns. and uses thereof)
TT
     Transforming growth factors
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (\beta-; osteogenic paste compns. and uses thereof)
     Microglobulins
IT
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (β-microglobulins; osteogenic paste compns. and uses thereof)
     56-81-5, Glycerol, biological studies 1306-06-5, Hydroxyapatite
IT
     7758-87-4, Tricalcium phosphate 61912-98-9, Insulin like growth factor
     62031-54-3, Fqf
     RL: DEV (Device component use); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
     USES (Uses)
        (osteogenic paste compns. and uses thereof)
REFERENCE COUNT:
                         9
                               THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L38 ANSWER 5 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER:
                         1999:139773 HCAPLUS
DOCUMENT NUMBER:
                         130:200953
TITLE:
                         A method of crosslinking collagen
                         -based material and bioprosthetic devices produced
INVENTOR(S):
                         Hendriks, Marc; Verhoeven, Michel; Cahalan, Patrick
                         T.; Torrianni, Mark W.; Fouache, Benedicte;
                         Cahalan, Linda
PATENT ASSIGNEE(S):
                         Medtronic, Inc., USA
SOURCE:
                         Eur. Pat. Appl., 26 pp.
                         CODEN: EPXXDW
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
    PATENT NO.
                        KIND
                                DATE
                                          APPLICATION NO.
                                                                  DATE
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     EP 897942
                                19990224 EP 1998-306595
                         A1
                                                                   19980818
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20040310
     EP 897942
                          В1
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO
     US 6166184
                                             US 1997-912778
                          Α
                                20001226
                                                                    19970818
                                             US 1997-912778
PRIORITY APPLN. INFO.:
                                                                 A 19970818
     Methods of crosslinking collagen-based material having
     collagen amine groups and collagen carboxyl groups are
     provided. The methods comprise blocking at least a portion of the
     collagen amine groups with a blocking agent to form blocked amine
     groups; contacting the collagen-based material having the
     blocked amine groups with a polyfunctional spacer; and activating at least
     a portion of the collagen carboxyl groups after blocking at
     least a portion of the collagen amine groups, wherein the
     polyfunctional spacer crosslinks the collagen-based
     material and wherein said contacting step may be effected before or after
     said activating step. Bioprosthetic devices made from these
     crosslinked collagen-based materials are also provided.
     Crosslinking involving the JEFFAMINE spacers shows the fastest
     rehydration, whereas glutaraldehyde crosslinking tends to be a
     bit slower. The highly hydrophilic crosslinked collagen
     -derived materials promote infiltration and diffusion of tissue fluid
     through the material matrix, providing supply of oxygen, nutritive
     substances, electrolytes and drainage of metabolites. Also, ingrowth of
     capillary blood vessels and cells is promoted, 25 and consequently the
     healing response is improved. In addition, hydrophilicity improves the blood
     compatibility of the material. Collagen samples
     crosslinked according to the method of the invention involving the
     Jeffamine D230 spacer had a cell growth inhibition of 25%, while cells
     with a deviant morphol. were not observed
IC
     ICM C08H001-06
     ICS A61L027-00
CC
     63-7 (Pharmaceuticals)
     Section cross-reference(s): 45
ST
     crosslinking collagen bioprosthetic device manuf
TT
     Acylation
        (agents; crosslinking collagen-based material for
        bioprosthetic devices manufacture)
IT
     Heart
        (aortic valve; crosslinking collagen-based material
        for bioprosthetic devices manufacture)
IT
     Collagens, biological studies
     RL: DEV (Device component use); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
        (crosslinked; crosslinking collagen-based
        material for bioprosthetic devices manufacture)
TT
     Biocompatibility
     Calcification
       Crosslinking
     Transplant and Transplantation
        (crosslinking collagen-based material for
        bioprosthetic devices manufacture)
IT
     Aldehydes, reactions
     Azides
     Ketones, reactions
     RL: NUU (Other use, unclassified); RCT (Reactant); RACT (Reactant or
     reagent); USES (Uses)
        (crosslinking collagen-based material for
        bioprosthetic devices manufacture)
IT
     Collagens, biological studies
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RL: RCT (Reactant); THU (Therapeutic use); BIOL (Biological study); RACT
     (Reactant or reagent); USES (Uses)
        (crosslinking collagen-based material for
        bioprosthetic devices manufacture)
     7732-18-5, Water, processes
IT
     RL: PEP (Physical, engineering or chemical process); PROC (Process)
        (absorption; crosslinking collagen-based material
        for bioprosthetic devices manufacture)
IT
     1122-58-3, 4-Dimethylaminopyridine 2592-95-2, N-Hydroxybenzotriazole
     6066-82-6, N-Hydroxysuccinimide
                                     39743-84-5
     RL: NUU (Other use, unclassified); USES (Uses)
        (crosslinking collagen-based material for
        bioprosthetic devices manufacture)
IT
     66-25-1, Hexanal 111-30-8, Glutaraldehyde 123-38-6, Propanal,
                123-72-8, Butanal 420-04-2, Cyanamide 530-62-1,
     reactions
     1,1'-Carbonyldiimidazole 538-75-0, N,N'-Dicyclohexylcarbodiimide
     616-02-4, Citraconic anhydride 693-13-0, N,N'-Diisopropylcarbodiimide 830-03-5, p-Nitrophenyl acetate 1865-01-6, p-Nitrophenyl formate
     2466-76-4, 1-Acetylimidazole 2491-17-0 2635-84-9, p-Nitrophenyl
     butyrate 6066-82-6D, N-Hydroxysuccinimide, esters 9046-10-0, Jeffamine
            14464-29-0, N-Hydroxysuccinimidyl acetate 16357-59-8,
     D 230
     2-Ethoxy-1-ethoxycarbonyl-1,2-dihydroquinoline 25952-53-8,
     1-Ethyl-3-(3-dimethylaminopropyl)carbodiimide hydrochloride
                                                                   30364-55-7
     74124-79-1, N,N'-Disuccinimidyl carbonate 94820-31-2 152305-87-8
     RL: NUU (Other use, unclassified); RCT (Reactant); RACT (Reactant or
     reagent); USES (Uses)
        (crosslinking collagen-based material for
        bioprosthetic devices manufacture)
IT
     74-94-2, Dimethylamine borane 75-22-9, Trimethylamine borane
                                                                      4856-95-5
     16940-66-2, Sodium borohydride 25895-60-7, Sodium cyanoborohydride
     65605-36-9
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (crosslinking collagen-based material for
        bioprosthetic devices manufacture)
REFERENCE COUNT:
                               THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS
                         6
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L38 ANSWER 6 OF 6 HCAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER:
                        1999:7871 HCAPLUS
DOCUMENT NUMBER:
                        130:57274
TITLE:
                        Bone graft composites and spacers
INVENTOR(S):
                        McKay, William F.
PATENT ASSIGNEE(S):
                        SDGI Holdings, Inc., USA
SOURCE:
                        PCT Int. Appl., 56 pp.
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
LANGUAGE:
                        English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
                                         APPLICATION NO.
     PATENT NO.
                        KIND
                               DATE
                              19981217 WO 1998-US11611 19980611
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     WO 9856433
                         A1
        W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
            DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG,
             KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
             NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
             UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,

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CM, GA, GN, ML, MR, NE, SN, TD, TG
     AU 9878185
                                 19981230
                                             AU 1998-78185
                                                                     19980611
                          A1
     AU 738218
                           B2
                                 20010913
     EP 988070
                                 20000329
                                             EP 1998-926323
                                                                     19980611
                          A1
     EP 988070
                                 20040915
                          В1
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
     JP 2002503992
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                                 20020205
                                             JP 1999-502905
                                                                     19980611
                                             AT 1998-926323
     AT 275981
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                                 20041015
                                                                     19980611
                                             US 1999-386560
     US 6261586
                          R1
                                 20010717
                                                                     19990831
PRIORITY APPLN. INFO.:
                                             US 1997-873276
                                                                  A 19970611
                                             WO 1998-US11611
                                                                  W 19980611
AB
     A bone graft substitute including a composition of natural selectively
     deactivated bone material which has been processed to remove associated non-
     collagenous bone proteins, said bone material containing native
     collagen materials and naturally associated bone minerals and
     substantially free from native non-collagenous protein, and a
     therapeutically effective amount to stimulate bone growth of a bone growth
     factor in synergistic combination with said bone material. Spacers
     composed of the bone graft substitute composition and methods for using the
     spacers are also provided. A diaphysial cortical bone dowel was prepared as
     well as deactivated allograft and its composite with BMP-2 composite.
IC
     ICM A61L027-00
     63-7 (Pharmaceuticals)
CC
ST
     bone graft composite spacer
     Bone morphogenetic proteins
     RL: PEP (Physical, engineering or chemical process); THU (Therapeutic
     use); BIOL (Biological study); PROC (Process); USES (Uses)
        (2; bone graft composites and spacers)
IT
     Bone
        (artificial; bone graft composites and spacers)
IT
     Collagens, biological studies
     Proteins, general, biological studies
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PEP
     (Physical, engineering or chemical process); THU (Therapeutic use); BIOL
     (Biological study); OCCU (Occurrence); PROC (Process); USES (Uses)
        (bone graft composites and spacers)
TT
     Growth factors, animal
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PEP
     (Physical, engineering or chemical process); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PROC (Process); USES (Uses)
        (bone-derived; bone graft composites and spacers)
IT
     Transplant and Transplantation
     Transplant and Transplantation
        (bone; bone graft composites and spacers)
IT
     Prosthetic materials and Prosthetics
        (composites, implants; bone graft composites and spacers)
IT
     Bone
     Bone
        (transplant; bone graft composites and spacers)
     1306-06-5, Hydroxyapatite 7758-87-4, Tricalcium phosphate
IT
     RL: PEP (Physical, engineering or chemical process); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
        (bone graft composites and spacers)
REFERENCE COUNT:
                         5
                                THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
                                RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
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